DONALD VOET

University of Pennsylvania

JUDITH G. VOET

Swarthmore College

BIOCHEMISTRY

SECOND EDITION



JOHN WILEY & SONS, INC. New York Chichester Brisbane Toronto Singapore Acquisitions Editor: Nedah Rose Marketing Manager: Catherine Faduska

Designer: Madelyn Lesure

Manufacturing Manager: Susan Stetzer Photo Researcher: Hilary Newman Illustration Coordinator: Edward Starr Illustrators: J/B Woolsey Associates

Production Management: Pamela Kennedy Oborski and

Suzanne Ingrao

Cover and Part Opening Illustrations © Irving Geis
This book was set in 9.5/12 Times Roman by Progressive
Information Technologies and printed and bound by
Von Hoffman Press. The cover was printed by Lehigh
Press Lithographers. Color separations by Lehigh Press
Colortronics.

Cover Art: Two paintings of horse heart cytochrome c by Irving Geis in which the protein is illuminated by its single iron atom. On the front cover the hydrophilic side chains are drawn in green, and on the back cover the hydrophobic side chains are drawn in orange. The paintings are based on an X-ray structure by Richard Dickerson.

Recognizing the importance of preserving what has been written, it is a policy of John Wiley & Sons, Inc. to have books of enduring value published in the United States printed on acid-free paper, and we exert our best efforts to that end.

John Wiley & Sons, Inc., places great value on the environment and is actively involved in efforts to preserve it. Currently, paper of high enough quality to reproduce full-color art effectively contains a maximum of 10% recovered and recycled post-consumer fiber. Wherever possible, Wiley uses paper containing the maximum amount of recycled fibers. In addition, the paper in this book was manufactured by a mill whose forest management programs include sustained yield harvesting of its timberlands. Sustained yield harvesting principles ensure that the number of trees cut each year does not exceed the amount of new growth.



Copyright © 1995, by John Wiley & Sons, Inc.

All rights reserved. Published simultaneously in Canada.

Reproduction or translation of any part of this work beyond that permitted by Sections 107 and 108 of the 1976 United States Copyright Act without the permission of the copyright owner is unlawful. Requests for permission or further information should be addressed to the Permissions Department, John Wiley & Sons, Inc.

ISBN: 0-471-58651-X

Printed in the United States of America

10 9 8 7 6 5 4 3 2

1. LIPID CLASSIFICATION

Lipids (Greek: lipos, fat) are substances of biological origin that are soluble in organic solvents such as chloroform and methanol but are only sparingly soluble, if at all, in water. Hence, they are easily separated from other biological materials by extraction into organic solvents and may be further fractionated by such techniques as adsorption chromatography, thin layer chromatography, and reverse-phase chromatography (Section 5-3E). Fats, oils, certain vitamins and hormones, and most nonprotein membrane components are lipids. In this section, we discuss the structures and physical properties of the major classes of lipids.

A. Fatty Acids

Fatty acids are carboxylic acids with long-chain hydrocarbon side groups (Fig. 11-1). They are rarely free in nature but, rather, occur in esterified form as the major components of the various lipids described in this chapter. The more common biological fatty acids are listed in Table 11-1. In higher plants and animals, the predominant fatty acid residues are those of the C_{16} and C_{18} species palmitic, oleic, linoleic, and stearic acids. Fatty acids with <14 or >20 carbon atoms are uncommon. Most fatty acids have an even number of carbon atoms because they are usually biosynthesized by the concatenation of C_2 units (Section 23-4C). Over half of the fatty acid residues of plant and animal lipids are unsaturated (contain double bonds) and are often

Stearic acid Oleic acid Linoleic acid α-Linolenic acid

FIGURE 11-1. The structural formulas of some C_{18} fatty acids. The double bonds all have the cis configuration.

TABLE 11-1. THE COMMON BIOLOGICAL FATTY ACIDS

Symbol*	Common Name	Systematic Name	Structure	mp (°C)
Saturated	fatty acids			
12:0	Lauric acid	Dodecanoic acid	CH ₃ (CH ₂) ₁₀ COOH	44.2
14:0	Myristic acid	Tetradecanoic acid	CH ₃ (CH ₂) ₁₂ COOH	52:
16:0	Palmitic acid	Hexadecanoic acid	CH ₃ (CH ₂) ₁₄ COOH	63.1
18:0	Stearic acid	Octadecanoic acid	CH ₃ (CH ₂) ₁₆ COOH	69.6
20:0	Arachidic acid	Eicosanoic acid	CH ₃ (CH ₂) ₁₈ COOH	75.4%
22:0	Behenic acid	Docosanoic acid	CH ₃ (CH ₂) ₂₀ COOH	81
24:0	Lignoceric acid	Tetracosanoic acid	CH ₃ (CH ₂) ₂₂ COOH	84.2
Unsaturate	d fatty acids (all doubl	e bonds are cis)		
16:1	Palmitoleic acid	9-Hexadecenoic acid	CH ₃ (CH ₂) ₃ CH=CH(CH ₃) ₃ COOH	-0.5
18:1	Oleic acid	9-Octadecenoic acid	$CH_3(CH_2)_7CH = CH(CH_2)_7COOH$	13.4
18:2	Linoleic acid	9,12-Octadecadienoic acid	$CH_3(CH_2)_4(CH=CHCH_2)_2(CH_2)_6COOH$	- 9 ₩
18:3	α -Linolenic acid	9,12,15-Octadecatrienoic acid	$CH_3CH_2(CH = CHCH_2)_3(CH_2)_6COOH$	-17
18:3	y-Linolenic acid	6,9,12-Octadecatrienoic acid	$CH_3(CH_2)_4(CH=CHCH_2)_3(CH_2)_3COOH$	7.8
20:4	Arachidonic acid	5,8,11,14-Eicosatetraenoic acid	$CH_3(CH_2)_4(CH=CHCH_2)_4(CH_2)_3COOH$	- 49 .5 🌋
20:5	EPA	5,8,11,14,17-Eicosapentanoic acid	CH ₃ CH ₂ (CH=CHCH ₂) ₅ (CH ₂) ₂ COOH	-54 ⊗
24:1	Nervonic acid	15-Tetracosenoic acid	$CH_3(CH_2)_7CH = CH(CH_2)_{17}COOH$	39

Number of carbon atoms: Number of double bonds.

polyunsaturated (contain two or more double bonds). Bacterial fatty acids are rarely polyunsaturated but are commonly branched, hydroxylated, or contain cyclopropane rings. Unusual fatty acids also occur as components of the oils and waxes (esters of fatty acids and long-chain alcohols) produced by certain plants.

The Physical Properties of Fatty Acids Vary with Their Degree of Unsaturation

Table 11-1 indicates that the first double bond of an unsaturated fatty acid commonly occurs between its C9 and C10 atoms counting from the carboxyl C atom (a Δ^9 - or 9-double bond). In polyunsaturated fatty acids, the double bonds tend to occur at every third carbon atom towards the methyl terminus of the molecule (such as CH=CH-CH₂-CH=CH-). Double bonds in polyunsaturated fatty acids are almost never conjugated (as in—CH=CH—CH=CH—). Triple bonds rarely occur in fatty acids or any other compound of biological origin.

Saturated fatty acids are highly flexible molecules that can assume a wide range of conformations because there is relatively free rotation about each of their C-C bonds. Nevertheless, their fully extended conformation is that of minimum energy because this conformation has the least amount of steric interference between neighboring methylene groups. The melting points (mp) of saturated fatty acids, like those of most substances, increase with molecular mass (Table 11-1).

Fatty acid double bonds almost always have the cis configuration (Fig. 11-1). This puts a rigid 30° bend in the hydrocarbon chain of unsaturated fatty acids that interferes with their efficient packing to fill space. The consequent reduced van der Waals interactions cause fatty acid melting points to decrease with their degree of unsaturation (Table (1-1). Lipid fluidity likewise increases with the degree of ansaturation of their component fatty acid residues. This phenomenon, as we shall see in Section 11-3B, has important consequences for membrane properties.

B. Triacylglycerols

The fats and oils that occur in plants and animals consist largely of mixtures of triacylglycerols (also referred to as triglycerides or neutral fats). These nonpolar, water-insoluble substances are fatty acid triesters of glycerol:

Triacylglycerols function as energy reservoirs in animals and are therefore their most abundant class of lipids even though they are not components of biological membranes.

Triacylglycerols differ according to the identity and placement of their three fatty acid residues. The so-called simple triacylglycerols contain one type of fatty acid residue and are named accordingly. For example, tristearoylglycerol or tristearin contains three stearic acid residues, whereas trioleoylglycerol or triolein has three oleic acid residues. The more common mixed triacylglycerols contain two or three different types of fatty acid residues and are named according to their placement on the glycerol moiety.

1-Palmitoleoyl-2-linoleoyl-3-stearoyl-glycerol

Fats and oils (which differ only in that fats are solid and oils are liquid at room temperature) are complex mixtures of simple and mixed triacylglycerols whose fatty acid compositions vary with the organism that has produced them. Plant oils are usually richer in unsaturated fatty acid residues than are animal fats, as the lower melting points of oils imply.